

Appl. No. 09/250,056  
Suppl. Arndt. Dated September 15, 2005

**Amendments to the Claims:**

Please amend claims 1, 4, 5, 11-13, 34, 39, and 40 as follows. No new matter has been added to the application by way of the above amendments, which are entirely formal in nature.

This listing of claims will replace all prior versions, and listings, of claims in the application.

**Listing of Claims:**

Claim 1. (Currently amended) An isolated, synthesized or recombinant antibody that specifically binds to a c-erbB2 receptor, wherein said antibody specifically binds to ~~an epitope of the c-erbB2 receptor, which epitope binds to~~ the c-erbB2 epitope as bound by F5 (SEQ ID NO:1) or C1 (SEQ ID NO:2), and further wherein said antibody is an internalizing antibody.

Claim 2. (Cancelled)

Claim 3. (Previously presented) The antibody of claim 1, wherein said antibody comprises an amino acid sequence selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 2, SEQ ID NO: 1 having conservative substitutions, and SEQ ID NO: 2 having conservative substitutions.

Claim 4. (Currently amended): The antibody of claim 1, wherein said antibody shares at least 70% sequence identity with the amino acid sequence of SEQ ID NO: 1 or SEQ ID NO: 2 as determined using a BLAST algorithm and default parameters, and wherein said antibody has a binding affinity for c-erbB2<sub>1</sub> on cells<sub>1</sub> of at least 10<sup>-7</sup> M.

Claim 5. (Currently amended): The antibody of claim 1, wherein ~~an~~ the amino acid sequence of said antibody differs from the amino acid sequence of SEQ ID NO: 1 or SEQ ID NO: 2 by no more than 30 residues.

Claim 6-10 (Cancelled)

Claim 11. (Currently amended): The antibody of claim 1, wherein said antibody comprises at least three complementarity determining regions ~~(CDRs) selected from the group consisting of the complementarity determining regions of~~ from SEQ ID NO: 1, ~~and or comprises at least three complementarity determining regions of~~ from SEQ ID NO: 2.

Claim 12. (Currently amended): The antibody of claim 11, wherein said antibody comprises the three complementarity determining regions of the amino acid sequence of SEQ ID NO: 1.

Appl. No. 09/250,056  
Suppl. Amdt. Dated September 15, 2005

Claim 13. (Currently amended): The antibody of claim 11, wherein said antibody comprises the three complementarity determining regions of the amino acid sequence of SEQ ID NO: 2.

Claim 14. (Previously presented): The antibody of claim 1, wherein said antibody comprises the amino acid sequence of SEQ ID NO: 1.

Claim 15. (Previously presented): The antibody of claim 1, wherein said antibody comprises the amino acid sequence of SEQ ID NO: 2.

Claims 16-33 (Canceled).

Claim 34. (Currently amended): A chimeric molecule that specifically binds a cell bearing a ~~a-e-erbB-2~~ c-erbB2 receptor, said chimeric molecule comprising an effector attached to an antibody of claim 1.

Claim 35. (Original): The chimeric molecule of claim 34, wherein said effector is selected from the group consisting of a cytotoxin, a label, a radionuclide, a drug, a liposome, a ligand, and an antibody.

Claim 36. (Original): The chimeric molecule of claim 34, wherein said chimeric molecule is a fusion protein.

Claim 37. (Original): The chimeric molecule of claim 34, wherein said cell is a cancer cell.

Claim 38. (Original): The chimeric molecule of claim 37, wherein said cancer cell is a breast cancer cell.

Claim 39. (Currently amended): The chimeric molecule of claim 34, wherein said antibody shares at least 70% sequence identity with the amino acid sequence of SEQ ID NO: 1 or SEQ ID NO: 2 as determined by a BLAST algorithm using default parameters, and wherein said antibody has a binding affinity for c-erbB2 of at least  $10^{-5}$  M.

Claim 40. (Currently amended): The chimeric molecule of claim 34, wherein ~~an~~ the amino acid sequence of said antibody differs from the amino acid sequence of SEQ ID NO: 1 or SEQ ID NO: 2 by no more than 30 residues.

Claim 41-42 (Cancelled)

Claim 43. (Previously presented): The chimeric molecule of claim 34, wherein said antibody comprises the amino acid sequence of SEQ ID NO: 1.

Claim 44. (Previously presented): The chimeric molecule of claim 34, wherein said antibody comprises the amino acid sequence of SEQ ID NO: 2.

Claims 45-52 (Cancelled)

Appl. No. 09/250,056  
Suppl. Amdt. Dated September 15, 2005

Claim 53. (Previously presented): A composition comprising a pharmacological excipient and the antibody of claim 1.

Claim 54. (Previously presented): A composition comprising a pharmacological excipient and the chimeric molecule of claim 34.

Claim 55. (Previously presented) The antibody of claim 1, wherein said antibody has a binding affinity of at least  $10^{-5}$  M.

Claim 56. (Previously presented): The antibody of claim 1, wherein said antibody comprises a single chain antibody.

Claim 57. (Previously presented): The antibody of claim 1, wherein said antibody comprises a homodimer.

Claim 58. (Previously presented): The antibody of claim 1, wherein said antibody is coupled to a surface of a phage.

Claim 59. (Previously presented): The antibody of claim 1, wherein the antibody comprises a chemically synthesized polypeptide, prepared by:

attaching a C-terminal amino acid of the polypeptide to an insoluble support; and  
sequentially adding remaining amino acids of the polypeptide, thereby preparing the chemically synthesized polypeptide.

Claim 60. (Previously presented): The antibody of claim 1, wherein the antibody is prepared from a nucleic acid sequence encoding said antibody by expressing the nucleic acid sequence in a cell.

Claim 61. (Previously presented): The antibody of claim 60, wherein the nucleic acid sequence encoding said antibody comprises a chain shuffled mutant scFv gene in which a  $V_H$  or  $V_L$  gene from SEQ ID NO: 1 or SEQ ID NO: 2 has been replaced with a human  $V_H$  or  $V_L$  gene.

Claim 62. (Previously presented): The antibody of claim 60, wherein the nucleic acid sequence encoding said antibody comprises a member of a mutant antibody sequence library in which one or more partial or entire CDR sequences have been randomized.

Claim 63. (Previously presented): The antibody of claim 60, wherein the nucleic acid sequence encoding said antibody comprises a member of a mutant antibody sequence library in which the CDRs of the member sequences are diversified by site directed mutagenesis.

Appl. No. 09/250,056  
Suppl. Amdt. Dated September 15, 2005

Claim 64. (Previously presented): The antibody of claim 60, wherein the nucleic acid sequence encoding said antibody is prepared by optimizing the nucleic acid sequence to reflect codon preferences for an expression system.

Claim 65. (Previously presented): The chimeric molecule of claim 34, said chimeric molecule comprising multiple effectors attached to the antibody of claim 1.

Claim 66. (Previously presented): The chimeric molecule of claim 34, said chimeric molecule comprising multiple targeting moieties.

Claim 67. (Previously presented): An isolated, synthesized, recombinant or single chain antibody that binds to a c-ErbB2 receptor, wherein the binding of said antibody to said receptor is reduced in the presence of F5 (SEQ ID NO: 1) or C1 (SEQ ID NO: 2), and wherein said antibody is an internalizing antibody.